**Serial No.**: 09/500,555 **Filed**: February 9, 2000

## Pending claims

- 1. (amended) An array composition comprising:
  - a) a substrate with a surface comprising discrete sites;
  - b) a population of microspheres comprising at least a first and a second subpopulation, wherein each subpopulation comprises a bioactive agent; and
  - c) at least one fiducial,
  - wherein said microspheres are distributed on said surface.
- 2. An array composition according to claim 1 wherein each subpopulation comprises a unique optical signature.
- 3.(amended) An array composition according to claim 1 wherein each subpopulation comprises an identifier binding ligand that will bind a decoder binding ligand for identification and elucidation of the bioactive agent.
- 4. An array composition according to claim 1 wherein said substrate is a fiber optic bundle and said fiducial is a fiducial fiber.
- 5. An array composition according to claim 1 wherein said substrate is a fiber optic bundle, said array comprises at least three non-linear fiducial, and each of said fiducial is a fiducial fiber.
- 6. An array composition according to claim 5 wherein at least one of said fiducial fibers has a different shape from the others.
- 7. An array composition according to claim 1 wherein said fiducial is a defined edge of said substrate.
- 8. An array composition according to claim 1 wherein said fiducial is a fiducial bead.
- 9. An array composition according to claim 1 wherein said bioactive agents are nucleic acids.
- 10. An array composition according to claim 1 wherein said bioactive agents are proteins.
- 11. An array composition according to claim 1 further comprising a computer readable memory comprising:
  - a) computer code that receives a first data image; and
  - b) computer code that registers said first data image using said fiducial to generate a first registered data image.

Serial No.: 09/500,555 Filed: February 9, 2000 12. An array composition according to claim 11 wherein said computer readable memory further comprises: a) computer code that receives a second data image; b) computer code that registers said second data image using said fiducial to generate a second registered data image; and c) computer code that compares said first and said second data image. 18. A method of making an array composition comprising: a) forming a surface comprising individual sites on a substrate; b) distributing microspheres on said surface such that said individual sites contain microspheres, wherein said microspheres comprise at least a first and a second subpopulations each comprising a bioactive agent; and c) incorporating at least one fiducial onto said surface. 19. (amended) A method according to claim 18 wherein said subpopulations further comprise an identifier binding ligand that will bind a decoder binding ligand for identification and elucidation of the bioactive agent. 20. (amended) A method according to claim 18 wherein said subpopulations further comprise an optical signature for identification and elucidation of the bioactive agent. 21. A method according to claim 18 wherein said substrate is a fiber optic bundle and said fiducial is a fiducial fiber. 22. A method according to claim 18 wherein said substrate is a fiber optic bundle, said array comprises at least three non-linear fiducial, and each of said fiducial is a fiducial fiber. 23. A method according to claim 122 wherein at least one of said fiducial fibers has a different shape from the others. 24. A method according to claim 18 wherein said fiducial is a defined edge of said substrate. 25. A method according to claim 18 wherein said fiducial is a fiducial bead. 26. A method according to claim 18 wherein said bioactive agents are nucleic acids. 27. A method according to claim 18 wherein said bioactive agents are proteins. 28. (new) A composition according to claim 1, wherein said discrete sites are wells. 29. (new) A composition according to claim 1, wherein said microspheres are randomly distributed on said substrate. 30. (new) A method according to claim 18, wherein said discrete sites are wells. - 10 -

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31. (new) A method according to claim 18, wherein said microspheres are randomly distributed on said substrate.